

Claim Amendments and Complete Listing of the Claims

1. – 4. (Canceled)

5. (Currently amended) A 3-[(indole-3-yl) methylene]-2-indolinone compound having a substituent at the 1' position of the indole, where the substituent at the 1' position is selected from the group consisting of,

(a) alkyl that is ~~optionally~~ substituted with a monocyclic or bicyclic five, six, eight, nine, or ten membered heterocyclic ring, where the ring is optionally substituted with one or more halogen, aldehyde, or trihalomethyl substituents;

(b) five, six, eight, nine, or ten membered monocyclic or bicyclic heterocyclic ring, where the ring is optionally substituted with one or more halogen or trihalomethyl substituents;

(c) an aldehyde or ketone of formula $-\text{CO}-\text{R}_{12}$, where R_{12} is selected from the group consisting of hydrogen, ~~alkyl~~, and a five or six membered heterocyclic ring;

(d) a carboxylic acid of formula $-(\text{R}_{13})_n-\text{COOH}$ or ester of formula $-(\text{R}_{14})_m-\text{COO}-\text{R}_{15}$, where R_{13} , R_{14} , and R_{15} are independently selected from the group consisting of alkyl and a five or six membered heterocyclic ring and m and n are independently 0 or 1;

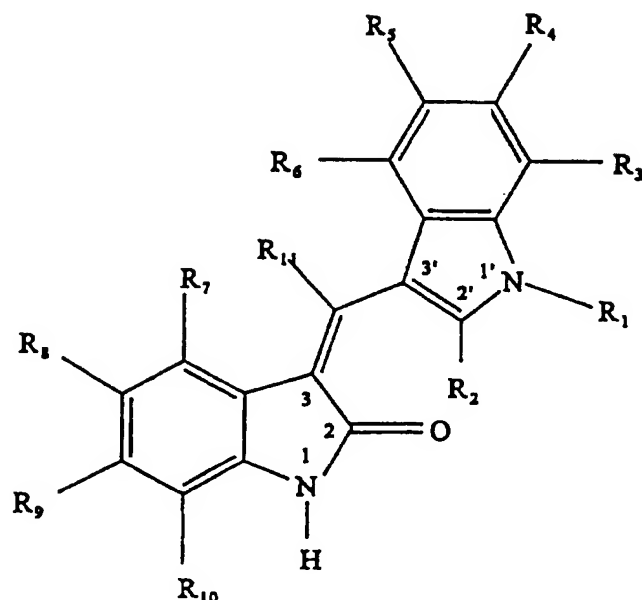
(e) a sulfone of formula $-(\text{SO}_2)-\text{R}_{16}$, ~~$-(\text{SO}_2)-\text{R}_{16}$~~ , where R_{16} is selected from the group consisting of alkyl and a five or six membered heterocyclic ring, where the ring is optionally substituted with an alkyl moiety;

(f) $-(\text{R}^{17})_n-(\text{indole-1-yl})$ or $-(\text{R}_{18})_m-\text{CHOH}-(\text{R}_{19})_p-(\text{indole-1-yl})$, where the indol moiety is optionally substituted with an aldehyde and R_{17} , R_{18} , and R_{19} are alkyl and m , n , and p are independently 0 or 1; and

(g) taken together with a 2' substituent of the indole ring forms a tricyclic moiety, where each ring in the tricyclic moiety is a five or six membered heterocyclic ring

or a salt, isomer, ester, amide, or prodrug thereof.

6. (Currently amended) The compound, salt, isomer, ~~metabolite~~, ester, amide, or prodrug of claim 5, wherein said compound has the formula,



where (a) R₁ is selected from the group consisting of,

- (i) alkyl that is ~~optionally~~ substituted with a monocyclic or bicyclic five, six, eight, nine, or ten membered heterocyclic ring, where the ring is optionally substituted with one or more halogen, aldehyde, or trihalomethyl substituents;
- (ii) five, six, eight, nine, or ten membered monocyclic or bicyclic heterocyclic ring, where the ring is optionally substituted with one or more halogen or trihalomethyl substituents;
- (iii) an aldehyde or ketone of formula -CO-R₁₂, where R₁₁ is selected from the group consisting of hydrogen, ~~alkyl, or~~ and a five or six membered heterocyclic ring;

(iv) a carboxylic acid of formula - $(R_{13})_n\text{-COOH}$ or ester of formula $\text{-}(R_{14})_m\text{-COO-}R_{15}$, ~~$\text{-}(R_{14})_m\text{-COO-}R_{15}$~~ , where R_{13} , R_{14} , and R_{15} are independently selected from the group consisting of alkyl or a five or six membered heterocyclic ring and n and m are independently 0 or 1;

(v) a sulfone of formula - $(\text{SO}_2)\text{-}R_{16}$, where R_{16} is selected from the group consisting of alkyl or a five or six membered heterocyclic ring, where the ring is optionally substituted with an alkyl moiety;

(vi) - $(R_{17})_n\text{-(indole-1-yl)}$ or $\text{-}(R_{18})_m\text{-CHOH-}(R_{19})_p\text{-(indole-1-yl)}$, ~~$\text{-}(R_{18})_m\text{-CHOH-}(R_{19})_p\text{-(indole-1-yl)}$~~ , where the indol moiety is optionally substituted with an aldehyde and R_{17} , R_{18} , and R_{19} are alkyl and n , m , and p are independently 0 or 1;

(vii) taken together with a 2' substituent of the indole ring forms a tricyclic moiety, where each ring in the tricyclic moiety is a five or six membered heterocyclic ring;

(b) R_2 , R_3 , R_4 , R_5 , and R_6 are selected from the group consisting of,

(i) hydrogen or alkyl that is optionally substituted with a monocyclic or bicyclic five, six, eight, nine, or ten membered heterocyclic ring, where the ring is optionally substituted with one or more halogen, aldehyde, or trihalomethyl substituents;

(ii) five, six, eight, nine, or ten membered monocyclic or bicyclic heterocyclic ring, where the ring is optionally substituted with one or more halogen or trihalomethyl substituents;

(iii) an aldehyde or ketone of formula $\text{-CO-}R_{20}$, where R_{20} is selected from the group consisting of hydrogen, alkyl, or a five or six membered heterocyclic ring;

(iv) a carboxylic acid of formula $\text{-}(R_{21})_n\text{-COOH}$ ~~$\text{-}(R_{21})_n\text{-COOH}$~~ or ester of formula - $(R_{22})\text{-COO-}R_{23}$, where R_{21} , R_{22} , and R_{23} are independently selected from the group

consisting of alkyl or a five or six membered heterocyclic ring and m and n are independently 0 or 1;

(v) halogen or an alcohol of formula $-(R_{24})_m-OH$, $(R_{24})_m-OH$ or an ether of formula $-(R_{24})_n-O-R_{25}$, where R_{24} and R_{25} are independently selected from the group consisting of alkyl and a five or six membered heterocyclic ring and m and n are independently 0 or 1;

(vi) $-NR_{26}R_{27}$, where R_{26} and R_{27} are independently selected from the group consisting of hydrogen, oxygen, alkyl, and a five or six membered heterocyclic ring; or $-NHCOR_{28}$, where R_{28} is selected from the group consisting of hydroxyl, alkyl, and a five or six membered heterocyclic ring, where the ring is optionally substituted with alkyl, halogen, carboxylate, or ester;

(vii) $-SO_2NR_{29}R_{30}$, $-SO_2NR_{29}R_{30}$, where R_{29} and R_{30} are selected from the group consisting of hydrogen, oxygen, alkyl, and a five or six membered heterocyclic ring;

(viii) any two of R_3 , R_4 , R_5 , or R_6 taken together form a bicyclic or tricyclic ~~heterocyclic~~ heterocyclic moiety fused to the six membered ring of the indole, where each ring in the multicyclic moiety is a five or six membered heterocyclic ring;

(c) R_7 , R_8 , R_9 , and R_{10} are independently selected from the group consisting of,

(i) hydrogen or alkyl that is optionally substituted with a monocyclic or bicyclic five, six, eight, nine, or ten membered heterocyclic ring, where the, ring is optionally substituted with one or more halogen, aldehyde, or trihalomethyl substituents;

(ii) five, six, eight, nine, or ten membered monocyclic or bicyclic heterocyclic ring, where the ring is optionally substituted with one or more halogen or trihalomethyl substituents;

(iii) an aldehyde or ketone of formula $-\text{CO}-\text{R}_{31}$, where R_{31} is selected from the group consisting of hydrogen, alkyl, or a five or six membered heterocyclic ring;

(iv) a carboxylic acid of formula $-(\text{R}_{32})_n-\text{COOH}$ ~~$-(\text{R}_{32})_m-\text{COOH}$~~ or ester of formula $-(\text{R}_{33})_m-\text{COO}-\text{R}_{34}$ ~~$-(\text{R}_{33})_m-\text{COO}-\text{R}_{34}$~~ , where R_{32} , R_{33} , and R_{34} and are independently selected from the group consisting of alkyl or a five or six membered heterocyclic ring and n and m are independently 0 or 1;

(v) halogen or an alcohol of formula $-(\text{R}_{35})_m-\text{OH}$ ~~$-(\text{R}_{35})_m-\text{OH}$~~ or an ether of formula $-(\text{R}_{35})_n-\text{O}-\text{R}_{36}$ ~~$-(\text{R}_{35})_n-\text{O}-\text{R}_{36}$~~ , where R_{35} and R_{36} are independently chosen from the group consisting of alkyl or a five or six membered heterocyclic ring and m and n are independently 0 or 1;

(vi) $-\text{NR}_{37}\text{R}_{38}$, where R_{37} and R_{38} are independently selected from the group consisting of hydrogen, oxygen, alkyl, and a five or six membered heterocyclic ring; or $-\text{NHCOR}_{39}$, where R_{39} is selected from the group consisting of hydroxyl, alkyl, and a five or six membered heterocyclic ring, where the ring is optionally substituted with alkyl, halogen, carboxylate, or ester;

(vii) $-\text{SO}_2\text{NR}_{29}\text{R}_{30}$ ~~$-\text{SO}_2\text{NR}_{29}\text{R}_{30}$~~ , where R_{40} and R_{41} are selected from the group consisting of hydrogen, oxygen, alkyl, and a five or six membered heterocyclic ring;

(viii) any two of R_7 , R_8 , R_9 , or R_{10} taken together form a bicyclic or tricyclic heterocyclic moiety fused to the six membered ring of the indole, where each ring in the multicyclic moiety is a five or six membered heterocyclic ring; and

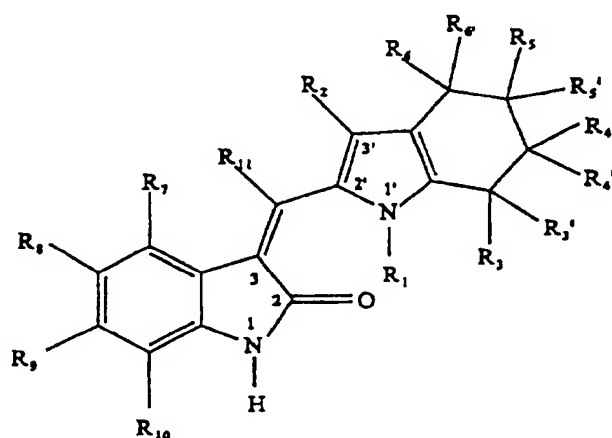
(d) R_{11} is hydrogen or alkyl;

provided that at least one of R_1 , R_2 , R_3 , R_4 , R_5 , R_6 , R_7 , R_8 , R_9 , or R_{10} is alkyl or provided that at least four of R_1 , R_2 , R_3 , R_4 , R_5 , or R_6 are not hydrogen.

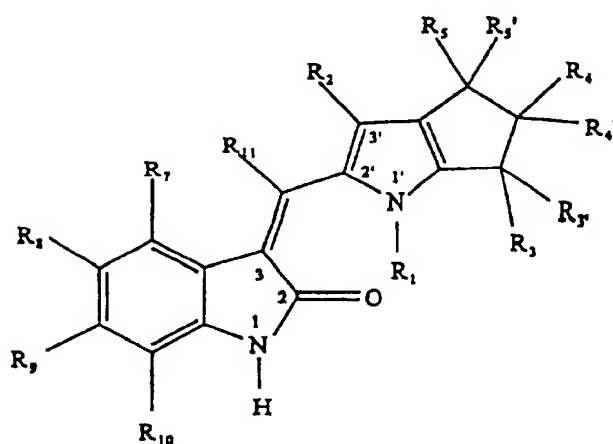
7. (Original) An optionally substituted 3-[(tetrahydroindole-2-yl) methylene] -2-indolinone or 3- [(cyclopentano-b-pyrrol-2-yl) methylene] -2-indolinone compound.

8. (Currently amended) The indolinone compound of claim 7 of formula XIX or XX,

XIX



XX



or a pharmaceutically acceptable salt, isomer, metabolite, ester, amide, or prodrug thereof where (a) R_1 is selected from the group consisting of,

- (i) alkyl that is optionally substituted with a monocyclic or bicyclic five, six, eight, nine, or ten membered heterocyclic ring, where the ring is optionally substituted with one or more halogen, or trihalomethyl substituents;
- (ii) five, six, eight, nine, or ten membered monocyclic or bicyclic heterocyclic ring, where the ring is optionally substituted with one or more halogen or trihalomethyl substituents;
- (iii) ketone of formula $-\text{CO}-R_{12}$, where R_{12} is selected from the group consisting of hydrogen, alkyl, or a five or six membered heterocyclic ring;
- (iv) a carboxylic acid of formula $-(R_{13})_n-\text{COOH}$ or ester of formula $-(R_{14})_m-\text{COO}-R_{15}$, where R_{13} , R_{14} , and R_{15} are independently selected from the group consisting of alkyl or a five or six membered heterocyclic ring and n and m are independently 0 or 1;
- (v) a sulfone of formula $-(\text{SO}_2)-R_{16}$, where R_{16} is selected from the group consisting of alkyl or a five or six membered heterocyclic ring, where the ring is optionally substituted with an alkyl moiety;
- (vi) $-(R_{17})_n-(\text{indole-1-yl})$ or $-(R_{18})_m-\text{CHOH}-(R_{19})_p-(\text{indole-1-yl})$, where the indole moiety is optionally substituted with an aldehyde and R_{17} , R_{18} , and R_{19} are alkyl and n , m , and p are independently 0 or 1;
- (vii) taken together with a 2' substituent of the indole ring forms a tricyclic moiety, where each ring in the tricyclic moiety is a five or six membered heterocyclic ring;

- (b) $R_2, R_3, R_3', R_4, R_4', R_5, R_5', R_6$ and R_6' are selected from the group consisting of,
- (i) hydrogen;
- (ii) alkyl that is optionally substituted with a monocyclic or bicyclic five, six, eight, nine, or ten membered heterocyclic ring, where the ring is optionally substituted with one or more halogen, or trihalomethyl substituents;
- (iii) five, six, eight, nine, or ten membered monocyclic or bicyclic heterocyclic ring, where the ring is optionally substituted with one or more halogen or trihalomethyl substituents;
- (iv) ketone of formula $-\text{CO}-R_{20}$, where R_{20} is selected from the group consisting of hydrogen, alkyl, or a five or six membered heterocyclic ring;
- (v) a carboxylic acid of formula $-(R_{21})_n-\text{COOH}$ or ester of formula $-(R_{22})-\text{COO}-R_{23}$, where R_{21}, R_{22} , and R_{23} are independently selected from the group consisting of alkyl or a five or six membered heterocyclic ring and m and n are independently 0 or 1;
- (vi) halogen;
- (vii) an alcohol of formula $-(R_{24})_m-\text{OH}$ or an ether of formula $-(R_{24})_n-\text{O}-R_{25}$, where R_{24} and R_{25} are independently selected from the group consisting of alkyl and a five or six membered heterocyclic ring and m and n are independently 0 or 1;
- (viii) $-\text{NR}_{26}\text{R}_{27}$, where R_{26} and R_{27} are independently selected from the group consisting of hydrogen, oxygen, alkyl, and a five or six membered heterocyclic ring;
- (ix) $-\text{NHCOR}_{28}$, where R_{28} is selected from the group consisting of hydroxyl, alkyl, and a five or six membered heterocyclic ring, where the ring is optionally substituted with alkyl, halogen, carboxylate, or ester;

- (x) $-\text{SO}_2\text{NR}_{29}\text{R}_{30}$, $-\text{SO}_2\text{NR}_{29}\text{R}_{30}$, where R_{29} and R_{30} are selected from the group consisting of hydrogen, oxygen, alkyl, and a five or six membered heterocyclic ring;
- (xi) any two of R_3 , R_3' , R_4 , R_4' , R_5 , R_5' , R_6 , or R_6' taken together form a bicyclic or tricyclic heterocyclic moiety fused to the six membered ring of the indole, where each ring in the multicyclic moiety is a five or six membered heterocyclic ring;
- (c) R_7 , R_8 , R_9 , and R_{10} are independently selected from the group consisting of,
- (i) hydrogen;
- (ii) alkyl that is optionally substituted with a monocyclic or bicyclic five, six, eight, nine, or ten membered heterocyclic ring, where the ring is optionally substituted with one or more halogen, or trihalomethyl substituents;
- (iii) five, six, eight, nine, or ten membered monocyclic or bicyclic heterocyclic ring, where the ring is optionally substituted with one or more halogen or trihalomethyl substituents;
- (iv) ketone of formula $-\text{CO}-\text{R}_{31}$, where R_{31} , is selected from the group consisting of hydrogen, alkyl, or a five or six membered heterocyclic ring;
- (v) a carboxylic acid of formula $-(\text{R}_{32})_n-\text{COOH}$ or ester of formula $-(\text{R}_{33})_m-\text{COO}-\text{R}_{34}$, where R_{32} , R_{33} , and R_{34} and are independently selected from the group consisting of alkyl or a five or six membered heterocyclic ring and n and m are independently 0 or 1;
- (vi) halogen;
- (vii) an alcohol of formula $(\text{R}_{35})_m-\text{OH}$ or an ether of formula $-(\text{R}_{35})_n-\text{O}-\text{R}_{36}$, where R_{35} and R_{36} are independently chosen from the group consisting of alkyl or a five or six membered heterocyclic ring and m and n are independently 0 or 1;

(viii) $-NR_{37}R_{38}$, where R_{37} and R_{38} are independently selected from the group consisting of hydrogen, oxygen, alkyl, and a five or six membered heterocyclic ring;

(ix) $-NHCOR_{39}$, where R_{39} is selected from the group consisting of hydroxyl, alkyl, and a five or six membered heterocyclic ring, where the ring is optionally substituted with alkyl, halogen, carboxylate, or ester;

(x) $-\underline{SO_2NR_{40}R_{41}}$, $-\underline{SO_2NR_{40}R_{41}}$, where R_{40} and R_{41} are selected from the group consisting of hydrogen, oxygen, alkyl, and a five or six membered heterocyclic ring;

(xi) any two of R_7 , R_8 , R_9 , or R_{10} taken together form a bicyclic or tricyclic ~~heterocyclic~~ heterocyclic moiety fused to the six membered ring of the indole, where each ring in the multicyclic moiety is a five or six membered heterocyclic ring; and

(d) R_{11} is hydrogen or alkyl.

9. (Currently amended) An indolinone compound having a substituent at the 5 position of the oxindole ring, where the substituent at the 5 position of the oxindole ring is selected from the group consisting of

(a) alkyl that is optionally substituted with a monocyclic or bicyclic five, six, eight, nine, or ten membered heterocyclic ring, where the ring is optionally substituted with one or more halogen, or trihalomethyl substituents;

(b) five, six, eight, nine, or ten membered monocyclic or bicyclic heterocyclic ring, where the ring is optionally substituted with one or more halogen or trihalomethyl substituents;

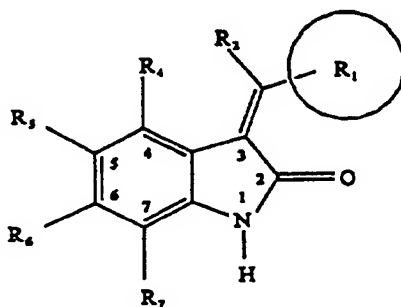
(c) a ketone of formula $-\text{CO}-R_{10}$, where R_{10} is selected from the group consisting of hydrogen, alkyl, or a five or six membered heterocyclic ring;

(d) a carboxylic acid of formula $-(R_{11})_n-\text{COOH}$ ~~$-(R_{11})_n-\text{COOH}$~~ or ester of formula $-(R_{12})-\text{COO}-R_{13}$, where R_{11} , R_{12} , and R_{13} and are independently selected from the group

consisting of alkyl or a five or six membered heterocyclic ring and m and n are independently 0 or 1;

- (e) halogen;
- (f) an alcohol of formula $-(R_{14})_m-OH$ or an ether of formula $-(R_{14})_n-O-R_{15}$, where R_{14} and R_{15} are independently selected from the group consisting of alkyl and a five or six membered heterocyclic ring and m and n are independently 0 or 1;
- (g) $-NR_{16}R_{17}$, where R_{16} and R_{17} are independently selected from the group consisting of hydrogen, alkyl, and a five or six membered heterocyclic ring;
- (h) $-NHCOR_{18}$, where R_{18} is selected from the group consisting of alkyl, and a five or six membered heterocyclic ring, where the ring is optionally substituted with alkyl, halogen, carboxylate, or ester;
- (i) $-SO_2NR_{19}R_{20}$, where R_{19} and R_{20} are selected from the group consisting of hydrogen, alkyl, and a five or six membered heterocyclic ring;
- (j) any two of R_4 , R_5 , R_6 , or R_7 taken together form a bicyclic or tricyclic heterocyclic moiety fused to the six membered ring of the oxindole, where each ring in the multicyclic moiety is a five or six membered heterocyclic ring.

10. (Currently amended) The compound of claim 9 of the following formula,



where (a) R_5 is selected from the group consisting of,

(i) alkyl that is optionally substituted with a monocyclic or bicyclic five, six, eight, nine, or ten membered heterocyclic ring, where the ring is optionally substituted with one or more halogen, or trihalomethyl substituents;

(ii) five, six, eight, nine, or ten membered monocyclic or bicyclic heterocyclic ring, where the ring is optionally substituted with one or more halogen or trihalomethyl substituents;

(iii) a ketone of formula $-\text{CO}-R_{10}$, where R_{10} is selected from the group consisting of hydrogen, alkyl, or a five or six membered heterocyclic ring;

(iv) a carboxylic acid of formula $-(R_{11})_n-\text{COOH}$ ~~$-(R_{11})_n-\text{COOH}$~~ or ester of formula $-(R_{12})-\text{COO}-R_{13}$, where R_{11} , R_{12} , and R_{13} are independently selected from the group consisting of alkyl or a five or six membered heterocyclic ring and m and n are independently 0 or 1;

(v) halogen;

(vi) an alcohol of formula $-(R_{14})_m-\text{OH}$ ~~$-(R_{14})_m-\text{OH}$~~ or an ether of formula $-(R_{14})_n-\text{O}-R_{15}$, where R_{14} and R_{15} are independently selected from the group consisting of alkyl and a five or six membered heterocyclic ring and m and n are independently 0 or 1;

(vii) $-\text{NR}_{16}\text{R}_{17}$, where R_{16} and R_{17} are independently selected from the group consisting of hydrogen, alkyl, and a five or six membered heterocyclic ring;

(viii) $-\text{NHCOR}_{18}$, where R_{18} is selected from the group consisting of alkyl, and a five or six membered heterocyclic ring, where the ring is optionally substituted with alkyl, halogen, carboxylate, or ester;

(ix) $-\text{SO}_2\text{NR}_{19}\text{R}_{20}$, where R_{19} and R_{20} are selected from the group consisting of hydrogen, alkyl, and a five or six membered heterocyclic ring;

(x) any two of R_4 , R_5 , R_6 , or R_7 taken together form a bicyclic or tricyclic heterocyclic ~~heterocyclic~~ moiety fused to the six membered ring of the oxindole, where each ring in the multicyclic moiety is a five or six membered heterocyclic ring;

(b) R_1 is selected from the group consisting of a five, six, eight, nine, and ten membered monocyclic or bicyclic heterocyclic ring, where the ring is optionally substituted with one or more substituents selected from the group consisting of

(i) hydrogen and alkyl that is optionally substituted with a monocyclic or bicyclic five, six, eight, nine, or ten membered heterocyclic ring, where the ring is optionally substituted with one or more halogen, or trihalomethyl substituents;

(ii) five, six, eight, nine, or ten membered monocyclic or bicyclic heterocyclic ring, where the ring is optionally substituted with one or more halogen or trihalomethyl substituents;

(iii) a ketone of formula $-\text{CO}-\text{R}_{21}$, where R_{21} is selected from the group consisting of hydrogen, alkyl, or a five or six membered heterocyclic ring;

(iv) a carboxylic acid of formula $-(\text{R}_{22})_m-\text{COOH}$ ~~$-(\text{R}_{22})_n-\text{COOH}$~~ or ester of formula $-(\text{R}_{23})-\text{COO}-\text{R}_{24}$, where R_{22} , R_{23} , and R_{24} are independently selected from the group consisting of alkyl or a five or six membered heterocyclic ring and m and n are independently 0 or 1;

(v) halogen;

(vi) an alcohol of formula $-(R_{25})_m-OH$ ~~$(R_{25})_m-OH$~~ or an ether of formula $-(R_{25})_n-O-$
 ~~R_{26}~~ , ~~$(R_{25})_n-O-R_{26}$~~ where R_{25} and R_{26} are independently selected from the group consisting of
 alkyl and a five or six membered heterocyclic ring and m and n are independently 0 or 1;

(vii) $-NR_{27}R_{28}$, where R_{27} and R_{28} are independently selected from the group consisting
 of hydrogen, alkyl, and a five or six membered heterocyclic ring;

(viii) $-NHCOR_{29}$, where R_{29} is selected from the group consisting of alkyl, and a five or
 six membered heterocyclic ring, where the ring is optionally substituted with alkyl, halogen,
 carboxylate, or ester;

(ix) $-SO_2NR_{30}R_{31}$, where R_{30} and R_{31} are selected from the group consisting of
 hydrogen, alkyl, and a five or six membered heterocyclic ring;

(c) R_4 , R_6 , and R_7 are independently selected from the group consisting of,

(i) hydrogen and alkyl that is optionally substituted with a monocyclic or bicyclic
 five, six, eight, nine, or ten membered heterocyclic ring, where the ring is optionally substituted
 with one or more halogen, or trihalomethyl substituents;

(ii) five, six, eight, nine, or ten membered monocyclic or bicyclic heterocyclic ring,
 where the ring is optionally substituted with one or more halogen or trihalomethyl substituents;

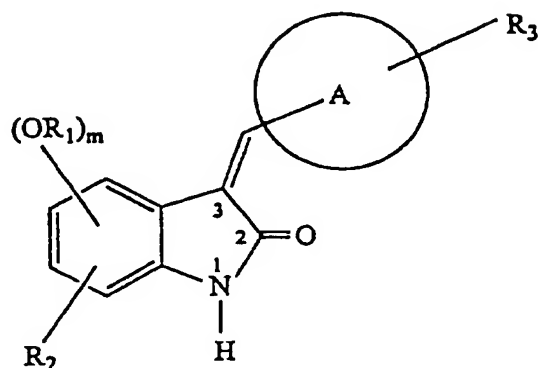
(iii) a ketone of formula $-CO-R_{32}$, where R_{32} is selected from the group consisting of
 hydrogen, alkyl, or a five or six membered heterocyclic ring;

(iv) a carboxylic acid of formula $-(R_{33})_n-COOH$ ~~$(R_{33})_n-COOH$~~ or ester of formula $-$
 $(R_{34})-COO-R_{35}$, where R_{33} , R_{34} and R_{35} are independently selected from the group consisting
 of alkyl or a five or six membered heterocyclic ring and m and n are independently 0 or 1;

(v) halogen;

- (vi) an alcohol of formula $-(R_{36})_m-OH$ ~~$(R_{36})_m-OH$~~ or an ether of formula $-(R_{36})_n-O-R_{37}$, where R_{36} and R_{37} are independently selected from the group consisting of alkyl and a five or six membered heterocyclic ring and m and n are independently 0 or 1;
- (vii) $-NR_{38}R_{39}$, where R_{38} and R_{39} are independently selected from the group consisting of hydrogen, alkyl, and a five or six membered heterocyclic ring;
- (viii) $-NHCOR_{40}$, where R_{40} is selected from the group consisting of alkyl, and a five or six membered heterocyclic ring, where the ring is optionally substituted with alkyl, halogen, carboxylate, or ester;
- (ix) $-SO_2NR_{41}R_{42}$, where R_{41} and R_{42} are selected from the group consisting of hydrogen, alkyl, and a five or six membered heterocyclic ring; and
- (d) R_2 is hydrogen or alkyl.

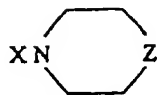
11. (Original) A compound having formula XXI, wherein:



XXI

- (a) A is a five or six membered ring comprised of atoms selected from the group consisting of oxygen, carbon, sulfur and nitrogen;
- (b) m is zero, 1, or 2;
- (c) R₁ is hydrogen, C₁-C₆ alkyl or C₂-C₆ alkanoyl;
- (d) one of R₂ and R₃ independently is hydrogen and the other is a substituent selected from:
- (1) a C₁-C₆ alkyl group substituted by 1, 2 or 3 hydroxy groups;
 - (2) SO₃R₄ in which R₄ is hydrogen or C₁-C₆ alkyl unsubstituted or substituted by 1, 2 or 3 hydroxy groups;
 - (3) SO₂NHR₅ in which R₅ is as R₄ defined above or a -(CH₂)_n-N(C₁-C₆ alkyl)₂ group in which n is 2 or 3;
 - (4) COOR₆ in which R₆ is C₁-C₆ alkyl unsubstituted or substituted by phenyl or by 1, 2 or 3 hydroxy groups or phenyl;
 - (5) CONHR₇ in which R₇ is hydrogen, phenyl or C₁-C₆ alkyl substituted by 1, 2 or 3 hydroxy groups or by phenyl;
 - (6) NHSO₂R₈ in which R₈ is C₁-C₆ alkyl or phenyl unsubstituted or substituted by halogen or by C₁-C₄ alkyl;
 - (7) N(R₉)₂, NHR₉ or OR₉ wherein R₉ is C₂-C₆ alkyl substituted by 1, 2 or 3 hydroxy groups;
 - (8) NHCOR₁₀, OOCR₁₀ or CH₂OOCR₁₀ in which R₁₀ is C₁-C₆ alkyl substituted by 1, 2 or 3 hydroxy groups;

(9) NHCONH_2 ; $\text{NH-C(NH}_2\text{)=NH}$; $\text{C(NH}_2\text{)=NH}$; $\text{CH}_2\text{NHC(NH}_2\text{)=NH}$; CH_2NH_2 ;
 OPO(OH)_2 ; $\text{CH}_2\text{OPO(OH)}_2$; PO(OH)_2 ; or a



wherein X is selected from the group consisting of CH_2 , SO_2 , CO , or $\text{NHCO(CH}_2\text{)}_p$ in which p is 1, 2, or 3 and Z is CH_2 , O or N-R_{11} in which R_{11} is hydrogen or is as R_9 defined above.

12. (Withdrawn) A method of making an indolinone compound of any one of claims 5-11 comprising the steps of reaching an appropriate aldehyde and oxindol and separating the indolinone from the aldehyde and oxindol reactants.

13. (Original) A pharmaceutical composition comprising (i) a pharmaceutically acceptable carrier or excipient and (ii) a compound according to any one of claims 5-11.

14. – 17. (Canceled)